Two Thousand Years of Lead Poisoning

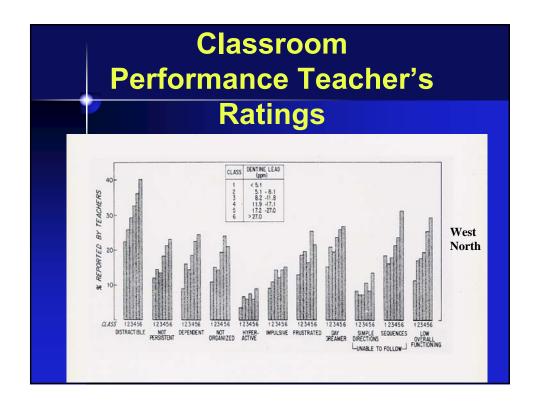
- •1st Century AD. Dioscerides: Lead makes the mind give way
- 1763. Benjamin Franklin: Lead in painters, printers and tinkers
- •1830 Charles Dickens: Uncommercial Traveler
- 1892.Brisbane Australia: Childhood lead poisoning described
- •1943. Randolph Byers: Long term effects
- ■1991. PHS Strategic Plan
- 1993: NAS Report

Five Phases of Lead Toxicology

- 1. There is no such thing as childhood lead poisoning. (1892)
- 2. If it doesn't kill you, it doesn't touch you
- 3. There are long term effects, but only in those with frank encephalpathy
- 4. Silent lead toxicity

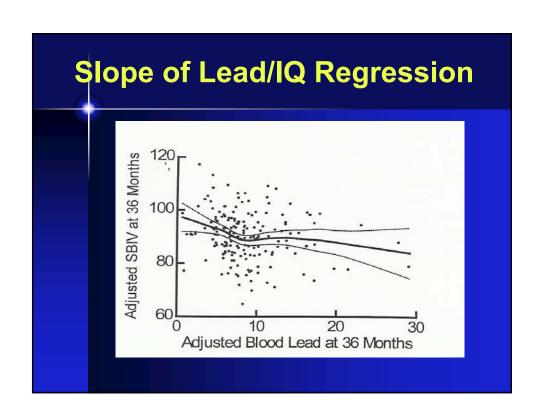
Flaws in Early Studies

- Inadequate exposure markers
- Screening or group measures of outcome
- Inadequate control of confounders
- Potential selection bias



Noncognitive effects

- Delinqency/social adjustment
- Attentional dysfunction
- Aging
- Cardiovascular effects/longevity



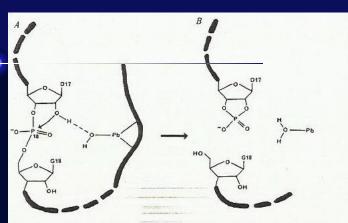
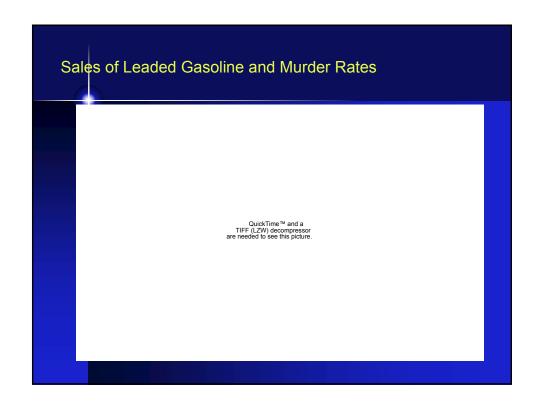
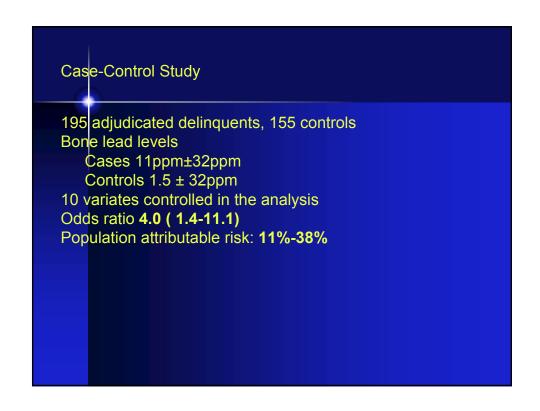


Fig. 2 Diagrammatic representation of the proposed mechanism for the sugar-phosphate backbone cleavage between residues D17 and G18 in monoclinic crystals of yeast tRNA^{Phe}. In A (before cleavage of the sugar-phosphate backbone) the Pb(II) moiety is depicted as being bound to, and positioned by, the tRNA^{Phe} molecule, while in B (after strand cleavage) the Pb(II) moiety is depicted as being free of the tRNA^{Phe} molecule.

Newer Studies

- Lanphear et al. Cognitive deficits associated with blood lead concentrations <10 microg/dL in US children and adolescents. Public Health Rep. 2000
- Canfield et al Intellectual impairment in children with blood lead concentrations below 10 microg per deciliter.NEJM 2003
- Bellinger and Needleman Intellectual impairment and blood lead levels NEJM 2003





Lessons Learned

- Require large samples: multicenter studies
- Multiple measures of exposure from birth onward of multiple toxins. Interaction effects
- Sensitive outcome measures: scheduled operative behavior, teachers' ratings, IQ
- Longterm follow up 40-50 years